

CLAIMS

1. A method for preparing peptides having selectively protected amines of
untargeted sites, comprising synthesizing the peptide by separately blocking branched
5 amines of targeted sites and branched amines of untargeted sites with either ivDde or
Mtt, and Boc, and protecting N^α-amine with Fmoc or Nsc.
2. The method in Claim 1, further comprising substituting the amine
protecting groups for amines of untargeted sites including N^α-amine with at least one
10 final amine protecting group selected from the group consisting of Fmoc, Nsc, Dde and
ivDde.
3. The method in Claim 1, further comprising substituting the amine
protecting group for amines of untargeted sites including N^α-amine with Boc.
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4. The method in any one of Claims 1 to 3, in which the peptide synthesis is
performed by solid phase synthesis.
5. The method in any one of Claims 1 to 3, in which the peptide sequence is
20 divided into at least two fragments, the fragments are synthesized separately, and then
the fragments are condensed to form the peptide.
6. Peptides having selectively protected amines of untargeted sites prepared
by the method in any one of Claims 1 to 5.

7. The peptides in Claim 6, in which said peptide is calcitonin or GRF(1-29).

8. A method for preparing specifically conjugated PEG-peptide in which PEG
5 is specifically conjugated to amines of targeted sites, comprising (1) a step of reacting
the peptide in Claim 6 with activated PEG and (2) a step of removing the amine
protecting group of the compound obtained in the step (1) under acid-base deblocking
condition.

10 9. The method in Claim 8, further comprising a step of purifying the product
of the step (2).

10. The method in Claim 9, in which said purification step comprises
separating the product by ionic exchange chromatography, removing salt and then
15 drying.

11. The method in any one of Claims 8 to 10, in which said activated PEG is
linear or branched hydroxy- or methoxy- type alkylating or acylating PEG of molecular
weight in a range of 1,000 to 40,000.

20 12. The method in Claim 11, in which said activated PEG is at least one
selected from the group consisting of mono-methoxy poly(ethyleneglycol)succinimidyl
succinate, mono-methoxy poly(ethyleneglycol)succinimidyl propionate, mono-methoxy
poly(ethyleneglycol)succinimidyl carbonate, mono-methoxy poly(ethyleneglycol)

succinimidyl carbamate and mono-methoxy poly(ethyleneglycol) tresylate.